

## SEQUENCE LISTING

<110> University of Newcastle Upon Tyne

<120> Fusion Proteins

<130> P69705US0

<140> US 10/501,071

<141> 2005-02-14

<150> GB 0200689.8

<151> 2002-01-10

<160> 62

<170> PatentIn version 3.1

<210> 1

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Ala3-His6 tail

<400> 1

Ala Ala Ala His His His His His 1

<210> 2

<211> 25

<212> PRT

<213> Escherichia coli

<400> 2

Met Asn Met Lys Lys Leu Ala Thr Leu Val Ser Ala Val Ala Leu Ser  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

```
Ala Thr Val Ser Ala Asn Ala Met Ala 20 25
<210> 3
<211> 5
<212> PRT
<213> Artificial Sequence
<220>
<223> Cleavage site for enterokinase
<400> 3
Asp Asp Asp Lys
1 5
<210> 4
<211> 4
<212> PRT
<213> Artificial Sequence
<220>
<223> Cleavage site for thrombin
<400> 4
Leu Val Pro Arg
<210> 5
<211> 4
<212> PRT
<213> Artificial Sequence
<220>
<223> Cleavage site for factor Xa
<400>
       5
Ile Glu Gly Arg
```

<210> 6

```
<211> 4
```

<212> PRT

<213> Artificial Sequence

<220>

<223> 4xHis tag

<400> 6

His His His His

<210> 7

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> 5xHis tag

<400> 7

His His His His 1 5

<210> 8

<211> 6

<212> PRT

<213> Artificial Sequence

<220>

<223> 6xHis tag

<400> 8

His His His His His 1

<210> 9

<211> 7

<212> PRT

```
<213> Artificial Sequence
<220>
<223> 7xHis tag
<400> 9
His His His His His His 1
<210> 10
<211> 8
<212> PRT
<213> Artificial Sequence
<220>
<223> 8xHis tag
<400> 10
His His His His His His His 1 5
<210> 11
<211> 9
<212> PRT
<213> Artificial Sequence
<220>
<223> 9xHis tag
<400> 11
His His His His His His His 1 5
<210> 12
<211> 10
<212> PRT
<213> Artificial Sequence
```

```
<223> 10xHis tag
```

<400> 12

His His His His His His His His His 10

<210> 13

<211> 93

<212> PRT

<213> Escherichia coli

<400> 13

Asn Asn Gly Ala Ser Gly Ala Asp Ile Asn Asn Tyr Ala Gly Gln Ile  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Lys Ser Ala Ile Glu Ser Lys Phe Tyr Asp Ala Ser Ser Tyr Ala Gly
20 25 30

Lys Thr Cys Thr Leu Arg Ile Lys Leu Ala Pro Asp Gly Met Leu Leu 35 40 45

Asp Ile Lys Pro Glu Gly Gly Asp Pro Ala Leu Cys Gln Ala Ala Leu 50 60

Ala Ala Ala Lys Leu Ala Lys Ile Pro Lys Pro Pro Ser Gln Ala Val 65 70 75 80

Tyr Glu Val Phe Lys Asn Ala Pro Leu Asp Phe Lys Pro 85 90

<210> 14

<211> 348

<212> PRT

<213> Artificial Sequence

<220>

<223> TolA-BCL fusion protein

<400> 14

Met His His His His His Ser Ser Asn Asn Gly Ala Ser Gly Ala 1 5 10 15 Asp Ile Asn Asn Tyr Ala Gly Gln Ile Lys Ser Ala Ile Glu Ser Lys 20 25 30 Phe Tyr Asp Ala Ser Ser Tyr Ala Gly Lys Thr Cys Thr Leu Arg Ile 35 40 45 Lys Leu Ala Pro Asp Gly Met Leu Leu Asp Ile Lys Pro Glu Gly Gly 50 60 Asp Pro Ala Leu Cys Gln Ala Ala Leu Ala Ala Lys Leu Ala Lys 65 70 75 80 Ile Pro Lys Pro Pro Ser Gln Ala Val Tyr Glu Val Phe Lys Asn Ala 85 90 95 Pro Leu Asp Phe Lys Pro Gly Gly Gly Ser Gly Ser Leu Val Pro Arg  $100 \hspace{1cm} 105 \hspace{1cm} 110$ Gly Ser Arg Pro Ser Gln Ser Asn Arg Glu Leu Val Val Asp Phe Leu 115 120 125 Ser Tyr Lys Leu Ser Gln Lys Gly Tyr Ser Trp Ser Gln Phe Ser Asp 130 135 140 Val Glu Glu Asn Arg Thr Glu Ala Pro Glu Gly Thr Glu Ser Glu Met 145 150 155 160 Glu Thr Pro Ser Ala Ile Asn Gly Asn Pro Ser Trp His Leu Ala Asp 165 170 175 Ser Pro Ala Val Asn Gly Ala Thr Ala His Ser Ser Ser Leu Asp Ala 180 185 190 Arg Glu Val Ile Pro Met Ala Ala Val Lys Gln Ala Leu Arg Glu Ala 195 200 205 Gly Asp Glu Phe Glu Leu Arg Tyr Arg Arg Ala Phe Ser Asp Leu Thr 210 220 Ser Gln Leu His Ile Thr Pro Gly Thr Ala Tyr Gln Ser Phe Glu Gln 225 230 235 Val Val Asn Glu Leu Phe Arg Asp Gly Val Asn Trp Gly Arg Ile Val 245 250 255 Ala Phe Phe Ser Phe Gly Gly Ala Leu Cys Val Glu Ser Val Asp Lys 260 265 270 Glu Met Gln Val Leu Val Ser Arg Ile Ala Ala Trp Met Ala Thr Tyr

Leu Asn Asp His Leu Glu Pro Trp Ile Gln Glu Asn Gly Gly Trp Asp 290 295 300

Thr Phe Val Glu Leu Tyr Gly Asn Asn Ala Ala Ala Glu Ser Arg Lys 305 310 315 320

Gly Gln Glu Arg Phe Asn Arg Trp Phe Leu Thr Gly Met Thr Val Ala 325 330 335

Gly Val Val Leu Leu Gly Ser Leu Phe Ser Arg Lys 340

<210> 15

<211> 236

<212> PRT

<213> Artificial Sequence

<220>

<223> Tola-BCL fusion protein after thrombin cleavage

<400> 15

Gly Ser Arg Pro Ser Gln Ser Asn Arg Glu Leu Val Val Asp Phe Leu 1 10 15

Ser Tyr Lys Leu Ser Gln Lys Gly Tyr Ser Trp Ser Gln Phe Ser Asp 20 25 30

Val Glu Glu Asn Arg Thr Glu Ala Pro Glu Gly Thr Glu Ser Glu Met 35 40 45

Glu Thr Pro Ser Ala Ile Asn Gly Asn Pro Ser Trp His Leu Ala Asp  $50 \hspace{1cm} 55 \hspace{1cm} 60$ 

Ser Pro Ala Val Asn Gly Ala Thr Ala His Ser Ser Ser Leu Asp Ala 65 70 75 80

Arg Glu Val Ile Pro Met Ala Ala Val Lys Gln Ala Leu Arg Glu Ala 85 90 95

Gly Asp Glu Phe Glu Leu Arg Tyr Arg Arg Ala Phe Ser Asp Leu Thr 100 105 110

Ser Gln Leu His Ile Thr Pro Gly Thr Ala Tyr Gln Ser Phe Glu Gln 115 120 125 Val Val Asn Glu Leu Phe Arg Asp Gly Val Asn Trp Gly Arg Ile Val 130 140

Ala Phe Phe Şer Phe Gly Gly Ala Leu Cys Val Glu Ser Val Asp Lys 145 150 155 160

Glu Met Gln Val Leu Val Ser Arg Ile Ala Ala Trp Met Ala Thr Tyr 165 170 175

Leu Asn Asp His Leu Glu Pro Trp Ile Gln Glu Asn Gly Gly Trp Asp 180 185 190

Thr Phe Val Glu Leu Tyr Gly Asn Asn Ala Ala Glu Ser Arg Lys 195 200 205

Gly Gln Glu Arg Phe Asn Arg Trp Phe Leu Thr Gly Met Thr Val Ala 210 220

Gly Val Val Leu Leu Gly Ser Leu Phe Ser Arg Lys 235 235

<210> 16

<211> 115

<212> PRT

<213> Artificial Sequence

<220>

<223> Tagged TolAIII region of pTol vectors

<220>

<221> MISC\_FEATURE

<222> (107)..(111)

<223> Xaa residues represent cleavage sites DDDDK (SEQ ID NO: 3), LVPR (SEQ ID NO: 4; no Xaa at position 111) or IEGR (SEQ ID NO: 5; no Xaa at position 111)

<400> 16

Met His His His His His Ser Ser Asn Asn Gly Ala Ser Gly Ala
1 5 10 15

Asp Ile Asn Asn Tyr Ala Gly Gln Ile Lys Ser Ala Ile Glu Ser Lys 20 25 30

Phe Tyr Asp Ala Ser Ser Tyr Ala Gly Lys Thr Cys Thr Leu Arg Ile 35 40 45 Lys Leu Ala Pro Asp Gly Met Leu Leu Asp Ile Lys Pro Glu Gly Gly 50 . 55 60 Asp Pro Ala Leu Cys Gln Ala Ala Leu Ala Ala Lys Leu Ala Lys 65 70 75 80 Ile Pro Lys Pro Pro Ser Gln Ala Val Tyr Glu Val Phe Lys Asn Ala  $85 \hspace{1cm} 90 \hspace{1cm} 95$ Pro Leu Asp Phe Lys Pro Gly Gly Gly Ser Xaa Xaa Xaa Xaa Xaa Gly 100 105 110 Ser Gly Thr 115 <210> 17 <211> 8 <212> PRT <213> Artificial Sequence <220> <223> His6-Ser2 linker <400> 17 His His His His His Ser Ser 1 5 <210> 18 <211> 4 <212> PRT <213> Artificial Sequence <220> Short flexible polypeptide <223>

Gly Gly Gly Ser

```
<210> 19
<211> 51
<212> DNA
<213> Artifiçial Sequence
<220>
<223> Cleavage/cloning site of pTolE vector
<400> 19
ggtgggggat ctgatgatga cgataaagga tccggtacct gatgaacgcg t
                                                                     51
<210> 20
<211> 48
<212> DNA
<213> Artificial Sequence
<220>
<223> Cleavage/cloning site of pTolT vector
<400> 20
ggtgggggat ctctggttcc gcgcggatcc ggtacctgat gaacgcgt
                                                                     48
<210> 21
<211> 48
<212> DNA
<213> Artificial Sequence
<220>
<223> Cleavage/cloning site of pTolX vector
<400>
ggtgggggat ctattgaagg tcgcggatcc ggtacctgat gaacgcgt
                                                                     48
<210> 22
<211> 17
<212> PRT
<213> Artificial Sequence
```

<220>

```
<223> Cleavage/cloning site of pTolE vector
<220>
<221> MISC_FEATURE
<222> (14)..(15)
<223> Xaa represents stop codon site
<400> 22
Gly Gly Ser Asp Asp Asp Asp Lys Gly Ser Gly Thr Xaa Xaa Thr 1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15
Arg
<210> 23
<211> 16
<212> PRT
<213> Artificial Sequence
<220>
       Cleavage/cloning site of pTolT vector
<220>
<221> MISC_FEATURE
<222> (13)..(14)
<223> Xaa represents stop codon site
<400> 23
Gly Gly Ser Leu Val Pro Arg Gly Ser Gly Thr Xaa Xaa Thr Arg 1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15
<210> 24
<211>
       16
<212> PRT
<213> Artificial Sequence
```

<220>

```
<223> Cleavage/cloning site of pTolX vector
<220>
<221> MISC_FEATURE
<222> (13)..(14)
<223> Xaa represents stop codon site
<400> 24
Gly Gly Gly Ser Ile Glu Gly Arg Gly Ser Gly Thr Xaa Xaa Thr Arg 1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15
<210> 25
<211> 2
<212> PRT
<213> Artificial Sequence
<220>
<223> Gly-Ser tag
<400> 25
Gly Ser
<210> 26
<211> 4
<212> PRT
<213> Artificial Sequence
<220>
<223> Gly-Ser-Gly-Thr tag
<400> 26
Gly Ser Gly Thr
<210> 27
<211> 39
<212> DNA
```

<213> Artificial Sequence

<220>		•
<223>	Synthetic oligonucleotide	
<400> gatctg	27 atga tgacgataaa ggatccggta cctgatgaa	39
<210>	28	
<211>	39	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Synthetic oligonucleotide	
<400> cgcgtt	28 catc aggtaccgga tcctttatcg tcatcatca	39
<210>	29	
<211>	36	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Synthetic oligonucleotide	
<400> gatcta	29 ttga aggtcgcgga tccggtacct gatgaa	36
<210>	30	
<211>	36	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Synthetic oligonucleotide	
<400> cgcgtt	30 catc aggtaccgga tccgcgacct tcaata	36

•

•

```
<210> 31
<211>
       36
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 31
                                                                              36
gatctctggt tccgcgcgga tccggtacct gatgaa
<210> 32
<211>
       36
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 32
                                                                              36
cgcgttcatc aggtaccgga tccgcgcgga accaga
<210> 33
<211> 37
<212> PRT
<213> Escherichia coli
<400> 33
Asn Ser Asn Gly Trp Ser Trp Ser Asn Lys Pro His Lys Asn Asp Gly 1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15
Phe His Ser Asp Gly Ser Tyr His Ile Thr Phe His Gly Asp Asn Asn 20 25 30
Ser Lys Pro Lys Pro
35
<210> 34
<211>
       80
```

<212> PRT

<213> Escherichia coli

<400> 34

Asn Asn Ala Phe Gly Gly Gly Lys Asn Pro Gly Ile Gly Asn Thr Ser  $1 \hspace{1cm} 10 \hspace{1cm} 15$ 

Gly Ala Gly Ser Asn Gly Ser Ala Ser Ser Asn Arg Gly Asn Ser Asn 20 25 30

Gly Trp Ser Trp Ser Asn Lys Pro His Lys Asn Asp Gly Phe His Ser 35 40 45

Asp Gly Ser Tyr His Ile Thr Phe His Gly Asp Asn Asn Ser Lys Pro 50 60

Lys Pro Gly Gly Asn Ser Gly Asn Arg Gly Asn Asn Gly Asp Gly Ala 65 70 75 80

<210> 35

<211> 117

<212> PRT

<213> Escherichia coli

<400> 35

His Gly Asp Asn Asn Ser Lys Pro Lys Pro Gly Gly Asn Ser Gly Asn 1 5 10 15

Arg Gly Asn Asn Gly Asp Gly Ala Ser Ala Lys Val Gly Glu Ile Thr 20 25 30

Ile Thr Pro Asp Asn Ser Lys Pro Gly Arg Tyr Ile Ser Ser Asn Pro 35 40 45

Glu Tyr Ser Leu Leu Ala Lys Leu Ile Asp Ala Glu Ser Ile Lys Gly 50 60

Thr Glu Val Tyr Thr Phe His Thr Arg Lys Gly Gln Tyr Val Lys Val 65 70 75 80

Thr Val Pro Asp Ser Asn Ile Asp Lys Met Arg Val Asp Tyr Val Asn 85 90 95

Trp Lys Gly Pro Lys Tyr Asn Asn Lys Leu Val Lys Arg Phe Val Ser 100 105 110 Gln Phe Leu Leu Phe 115

<210> 36

<211> 124

<212> PRT

<213> Homo sapiens

<400> 36

Asn Leu Val Asn Phe His Arg Met Ile Lys Leu Thr Thr Gly Lys Glu
1 10 15

Ala Ala Leu Ser Tyr Gly Phe Tyr Gly Cys His Cys Gly Val Gly Gly 20 25 30

Arg Gly Ser Pro Lys Asp Ala Thr Asp Arg Cys Cys Val Thr His Asp 35 40 45

Cys Cys Tyr Lys Arg Leu Glu Lys Arg Gly Cys Gly Thr Lys Phe Leu 50 60

Ser Tyr Lys Phe Ser Asn Ser Gly Ser Arg Ile Thr Cys Ala Lys Gln 65 70 75 80

Asp Ser Cys Arg Ser Gln Leu Cys Glu Cys Asp Lys Ala Ala Ala Thr 85 90 95

Cys Phe Ala Arg Asn Lys Thr Thr Tyr Asn Lys Lys Tyr Gln Tyr Tyr 100 105 110

Ser Asn Lys His Cys Arg Gly Ser Thr Pro Arg Cys 115 120

<210> 37

<211> 179

<212> PRT

<213> Actinia equina

<400> 37

Ser Ala Asp Val Ala Gly Ala Val Ile Asp Gly Ala Ser Leu Ser Phe 1 10 15

Asp Ile Leu Lys Thr Val Leu Glu Ala Leu Gly Asn Val Lys Arg Lys 20 25 30

Ile Ala Val Gly Val Asp Asn Glu Ser Gly Lys Thr Trp Thr Ala Leu 35 . 40 45

Asn Thr Tyr Phe Arg Ser Gly Thr Ser Asp Ile Val Leu Pro His Lys 50 . 55 60

Val Pro His Gly Lys Ala Leu Leu Tyr Asn Gly Gln Lys Asp Arg Gly 65 70 75 80

Pro Val Ala Thr Gly Ala Val Gly Val Leu Ala Tyr Leu Met Ser Asp 85 90 95

Gly Asn Thr Leu Ala Val Leu Phe Ser Val Pro Tyr Asp Tyr Asn Trp  $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110$ 

Tyr Ser Asn Trp Trp Asn Val Arg Ile Tyr Lys Gly Lys Arg Arg Ala 115 120 125

Asp Gln Arg Met Tyr Glu Glu Leu Tyr Tyr Asn Leu Ser Pro Phe Arg 130 135 140

Gly Asp Asn Gly Trp His Thr Arg Asn Leu Gly Tyr Gly Leu Lys Ser 150 155 160

Arg Gly Phe Met Asn Ser Ser Gly His Ala Ile Leu Glu Ile His Val 165 170 175

Ser Lys Ala

<210> 38

<211> 191

<212> PRT

<213> Homo sapiens

<400> 38

Thr Gly Ala Gly Lys Thr Ser Leu Leu Met Met Ile Met Gly Glu Leu 1 5 10 15

Glu Pro Ser Glu Gly Lys Ile Lys His Ser Gly Arg Ile Ser Phe Cys 20 25 30

Ser Gln Phe Ser Trp Ile Met Pro Gly Thr Ile Lys Glu Asn Ile Ile 35 40 45

Phe Gly Val Ser Tyr Asp Glu Tyr Arg Tyr Arg Ser Val Ile Lys Ala 50 55 60

Cys Gln Leu Glu Glu Asp Ile Ser Lys Phe Ala Glu Lys Asp Asn Ile 65 70 75 80

Val Leu Gly Glu Gly Gly Ile Thr Leu Ser Gly Gly Gln Arg Ala Arg 85 90 95

Ile Ser Leu Ala Arg Ala Val Tyr Lys Asp Ala Asp Leu Tyr Leu Leu 100 105 110

Asp Ser Pro Phe Gly Tyr Leu Asp Val Leu Thr Glu Lys Glu Ile Phe 115 120 125

Glu Ser Cys Val Cys Lys Leu Met Ala Asn Lys Thr Arg Ile Leu Val 130 135 140

Thr Ser Lys Met Glu His Leu Lys Lys Ala Asp Lys Ile Leu Ile Leu 145 150 155 160

His Glu Gly Ser Ser Tyr Phe Tyr Gly Thr Phe Ser Glu Leu Gln Asn 165 170 175

Leu Gln Pro Asp Phe Ser Ser Lys Leu Met Gly Cys Asp Ser Phe  $180\,$   $185\,$  190

<210> 39

<211> 390

<212> PRT

<213> Homo sapiens

<400> 39

Lys Tyr Ile Glu His Phe Ser Lys Phe Ser Pro Ser Pro Leu Ser Met
1 5 10 15

Lys Gln Phe Leu Asp Phe Gly Ser Ser Asn Ala Cys Glu Lys Thr Ser 20 25 30

Phe Thr Phe Leu Arg Gln Glu Leu Pro Val Arg Leu Ala Asn Ile Met 35 40 . 45

Lys Glu Ile Asn Leu Leu Pro Asp Arg Val Leu Ser Thr Pro Ser Val

55

Gln Leu Val Gln Ser Trp Tyr Val Gln Ser Leu Leu Asp Ile Met Glu 65 70 75 80 Phe Leu Asp Lys Asp Pro Glu Asp His Arg Thr Leu Ser Gln Phe Thr 85 90 95 Asp Ala Leu Val Thr Ile Arg Asn Arg His Asn Asp Val Val Pro Thr 100 105 110 Met Ala Gln Gly Val Leu Glu Tyr Lys Asp Thr Tyr Gly Asp Asp Pro 115 120 125 Val Ser Asn Gln Asn Ile Gln Tyr Phe Leu Asp Arg Phe Tyr Leu Ser 130 135 140 Arg Ile Ser Ile Arg Met Leu Ile Asn Gln His Thr Leu Ile Phe Asp 145 150 155 160 Gly Ser Thr Asn Pro Ala His Pro Lys His Ile Gly Ser Ile Asp Pro 165 170 175 Asn Cys Asn Val Ser Glu Val Val Lys Asp Ala Tyr Asp Met Ala Lys 180 185 190 Leu Leu Cys Asp Lys Tyr Tyr Met Ala Ser Pro Asp Leu Glu Ile Gln 195 200 205 Glu Ile Asn Ala Ala Asn Ser Lys Gln Pro Ile His Met Val Tyr Val 210 215 220 Pro Ser His Leu Tyr His Met Leu Phe Glu Leu Phe Lys Asn Ala Met 225 230 235 240 Arg Ala Thr Val Glu Ser His Glu Ser Ser Leu Ile Leu Pro Pro Ile 245 250 255 Lys Val Met Val Ala Leu Gly Glu Glu Asp Leu Ser Ile Lys Met Ser 260 265 270 Asp Arg Gly Gly Val Pro Leu Arg Lys Ile Glu Arg Leu Phe Ser 275 280 285

Pro Leu Ala Gly Phe Gly Tyr Gly Leu Pro Ile Ser Arg Leu Tyr Ala 305 310 315 320

Tyr Met Tyr Ser Thr Ala Pro Thr Pro Gln Pro Gly Thr Gly Gly Thr 290 295 300 Lys Tyr Phe Gln Gly Asp Leu Gln Leu Phe Ser Met Glu Gly Phe Gly 325 330 335

Thr Asp Ala Val Ile Tyr Leu Lys Ala Leu Ser Thr Asp Ser Val Glu 340 345 350

Arg Leu Pro Val Tyr Asn Lys Ser Ala Trp Arg His Tyr Gln Thr Ile 355 360 365

Gln Glu Ala Gly Asp Trp Cys Val Pro Ser Thr Glu Pro Lys Asn Thr 370 375 380

Ser Thr Tyr Arg Val Ser 385 390

<210> 40

<211> 202

<212> PRT

<213> Bacillus licheniformis

<400> 40

Ser Phe Ser Glu Leu Asn Ile Asp Ala Phe Arg Phe Ile Asn Asp Leu 1 5 10 15

Gly Lys Glu Tyr Ser Met Leu Asn Pro Val Val Tyr Phe Leu Ala Glu 20 25 30

Tyr Met Met Tyr Phe Leu Ala Leu Gly Leu Val Val Tyr Trp Leu Thr 35 40 45

Arg Thr Thr Lys Asn Arg Leu Met Val Ile Tyr Ala Val Ile Ala Phe 50 55 60

Val Val Ala Glu Ile Leu Gly Lys Ile Met Gly Ser Leu His Ser Asn 65 70 75 80

Tyr Gln Pro Phe Ala Thr Leu Pro Asn Val Asn Lys Leu Ile Glu His 85 90 95

Glu Ile Asp Asn Ser Phe Pro Ser Asp His Thr Ile Leu Phe Phe Ser 100 105 110

Ile Gly Phe Leu Ile Phe Leu Phe His Lys Lys Thr Gly Trp Leu Trp 115 120 125

Leu Val Leu Ala Phe Ala Val Gly Ile Ser Arg Ile Trp Ser Gly Val 130 135 140

His Tyr Pro Leu Asp Val Ala Ala Gly Ala Leu Leu Gly Val Leu Ser 145 150 155 160

Ala Leu Phe Val Phe Trp Thr Ala Pro Lys Leu Ser Phe Ile His Gln
• 165 170 175

Met Leu Ser Leu Tyr Glu Lys Val Glu Gln Arg Ile Val Pro Ser Lys 180 185 190

Asn Lys Ser Asn Asp Lys Ser Lys Asn Phe 195 200

<210> 41

<211> 354

<212> PRT

<213> Homo sapiens

<400> 41

Gln Arg Ser Pro Leu Glu Lys Ala Ser Val Val Ser Lys Leu Phe Phe 1 5 10 15

Ser Trp Thr Arg Pro Ile Leu Arg Lys Gly Tyr Arg Gln Arg Leu Glu 20 25 30

Leu Ser Asp Ile Tyr Gln Ile Pro Ser Val Asp Ser Ala Asp Asn Leu 35 40 45

Ser Glu Lys Leu Glu Arg Glu Trp Asp Arg Glu Leu Ala Ser Lys Lys 50 55 60

Asn Pro Lys Leu Ile Asn Ala Leu Arg Arg Cys Phe Phe Trp Arg Phe 65 70 75 80

Met Phe Tyr Gly Ile Phe Leu Tyr Leu Gly Glu Val Thr Lys Ala Val 85 90 95

Gln Pro Leu Leu Gly Arg Ile Ile Ala Ser Tyr Asp Pro Asp Asn  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Lys Glu Glu Arg Ser Ile Ala Ile Tyr Leu Gly Ile Gly Leu Cys Leu 115 120 125 Leu Phe Ile Val Arg Thr Leu Leu Leu His Pro Ala Ile Phe Gly Leu 130 135 140

His His Ile Gly Met Gln Met Arg Ile Ala Met Phe Ser Leu Ile Tyr 145 150 155 160

Lys Lys Thr Leu Lys Leu Ser Ser Arg Val Leu Asp Lys Ile Ser Ile 165 170 175

Gly Gln Leu Val Ser Leu Leu Ser Asn Asn Leu Asn Lys Phe Asp Glu 180 185 190

Gly Leu Ala Leu Ala His Phe Val Trp Ile Ala Pro Leu Gln Val Ala 195 200 205

Leu Leu Met Gly Leu Ile Trp Glu Leu Leu Gln Ala Ser Ala Phe Cys 210 215 220

Gly Leu Gly Phe Leu Ile Val Leu Ala Leu Phe Gln Ala Gly Leu Gly 225 230 235 240

Arg Met Met Lys Tyr Arg Asp Gln Arg Ala Gly Lys Ile Ser Glu 245 250 255

Arg Leu Val Ile Thr Ser Glu Met Ile Glu Asn Ile Gln Ser Val Lys 260 265 270

Ala Tyr Cys Trp Glu Glu Ala Met Glu Lys Met Ile Glu Asn Leu Arg 275 280 285

Gln Thr Glu Leu Lys Leu Thr Arg Lys Ala Ala Tyr Val Arg Tyr Phe 290 295 300

Asn Ser Ser Ala Phe Phe Phe Ser Gly Phe Phe Val Val Phe Leu Ser 305 310 315 320

Val Leu Pro Tyr Ala Leu Ile Lys Gly Ile Ile Leu Arg Lys Ile Phe 325 330 335

Thr Thr Ile Ser Phe Cys Ile Val Leu Arg Met Ala Val Thr Arg Gln
340 345 350

Phe Pro

<210> 42

<211> 34

<212> DNA

```
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 42
                                                                      34
tttttggatc caattccaat ggatggtcat ggag
<210> 43
<211> 41
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 43
                                                                      41
aaggatccaa gcttcaaggt ttaggctttg aattattgtc c
<210> 44
<211> 36
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 44
                                                                      36
tttttggatc caatgctttt ggtggaggga aaaatc
<210> 45
<211> 19
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 45
                                                                      19
ctcagcggtg gcagcagcc
```

```
<210> 46
<211> 31
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 46
                                                                     31
cgcggatccc atggggacaa taattcaaag c
<210> 47
<211> 38
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
                                                                     38
ggcgaattca cgcgttaaaa taataatttc tggctcac
<210> 48
<211> 37
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 48
ccggggtacc aatttggtga atttccacag aatgatc
                                                                     37
<210> 49
<211> 35
<212> DNA
<213> Artificial Sequence
```

<220>			
<223>	Synthetic oligonucleotide		
<400> ggcgaa	49 ttca cgcgttagca acgaggggtg ctccc	35	
<210>	50		
<211>	27		
<212>	DNA		
<213>	Artificial Sequence		
<220>			
<223>	Synthetic oligonucleotide		
<400> cgcgga	50 tccg cagacgtggc tggcgcc	27	
<210>	51		
<211>	38		
<212>	DNA		
<213>	Artificial Sequence		
<220>			
<223>	Synthetic oligonucleotide		
<400> 51 ggcgaattca cgcgttaagc tttgctcacg tgagtttc 38			
<210>	52		
<211>	30		
<212>	DNA		
<213>	Artificial Sequence		
<220>			
<223>	Synthetic oligonucleotide		
<400> cgcgga	52 tcct ctaatggtga tgacagcctc	30	
<210>	53		

```
<211> 38
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 53
                                                                     38
ggcgaattca cgcgttagaa agaatcacat cccatgag
<210> 54
<211> 37
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
<400> 54
                                                                     37
ccggggtacc aagtacatag agcacttcag caagttc
<210> 55
<211> 37
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
                                                                     37
ggcgaattca cgcgttacgt gacgcggtac gtggtcg
<210> 56
<211> 31
<212> DNA
<213> Artificial Sequence
<220>
<223> Synthetic oligonucleotide
```

<pre>&lt;400&gt; 56 cgcggatcct tttcagaatt aaatattgat g . 31</pre>				
<210>	57			
<211>	36			
<212>	DNA 2			
<213>	Artificial Sequence			
<220>				
<223>	Synthetic oligonucleotide			
<400> ggcgaa	57 ttca cgcgttaaaa gttcttcgat ttatcg	36		
<210>	58			
<211>	25			
<212>	DNA			
<213>	Artificial Sequence			
<220>				
<223>	Synthetic oligonucleotide			
<400>	58 tccc agaggtcgcc tctgg	25		
cycyyu	teee agaggeegee teegg			
<210>	59			
<211>	35			
<212>	DNA			
<213>	Artificial Sequence			
<220>				
<223>	Synthetic oligonucleotide			
<400> 59 ggcgaattca cgcgttaggg aaattgccga gtgac 35				
ggcgaa	cee egegeenggg municipeegn gegne	- •		
<210>	60			
<211>	30			

<212> DNA

<213>	Artificial	Sequence
	,	2042000

<220>

41.4

<223> Synthetic oligonucleotide

<400> 60

ttttttaggc cttctcagag caaccgggag

30

28

<210> 61

<211> 28

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic oligonucleotide

<400> 61

ttttacgcgt tcatttccga ctgaagag

<210> 62

<211> 233

<212> PRT

<213> Homo sapiens

<400> 62

Met Ser Gln Ser Asn Arg Glu Leu Val Val Asp Phe Leu Ser Tyr Lys  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Leu Ser Gln Lys Gly Tyr Ser Trp Ser Gln Phe Ser Asp Val Glu Glu 20 25 30

Asn Arg Thr Glu Ala Pro Glu Gly Thr Glu Ser Glu Met Glu Thr Pro 35 40 45

Ser Ala Ile Asn Gly Asn Pro Ser Trp His Leu Ala Asp Ser Pro Ala 50 55 60

Val Asn Gly Ala Thr Ala His Ser Ser Ser Leu Asp Ala Arg Glu Val 65 70 75 80

Ile Pro Met Ala Ala Val Lys Gln Ala Leu Arg Glu Ala Gly Asp Glu 85 90 95

Phe Glu Leu Arg Tyr Arg Arg Ala Phe Ser Asp Leu Thr Ser Gln Leu  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

His Ile Thr Pro Gly Thr Ala Tyr Gln Ser Phe Glu Gln Val Val Asn 115 120 125

Glu Leu Phe Arg Asp Gly Val Asn Trp Gly Arg Ile Val Ala Phe Phe 130 40 140

Ser Phe Gly ©ly Ala Leu Cys Val Glu Ser Val Asp Lys Glu Met Gln 145 150 155 160

Val Leu Val Ser Arg Ile Ala Ala Trp Met Ala Thr Tyr Leu Asn Asp 165 170 175

His Leu Glu Pro Trp Ile Gln Glu Asn Gly Gly Trp Asp Thr Phe Val 180 185 190

Glu Leu Tyr Gly Asn Asn Ala Ala Ala Glu Ser Arg Lys Gly Gln Glu 195 200 205

Arg Phe Asn Arg Trp Phe Leu Thr Gly Met Thr Val Ala Gly Val Val 210 220

Leu Leu Gly Ser Leu Phe Ser Arg Lys 235